

10/11/2006 10566562a1.trn

Connecting via Winsock to STN

Welcome to STN International! Enter x:x

LOGINID:SSSPTA1626GMS

PASSWORD:

TERMINAL (ENTER 1, 2, 3, OR ?):2

\* \* \* \* \* Welcome to STN International \* \* \* \* \*

NEWS 1 Web Page URLs for STN Seminar Schedule - N. America  
NEWS 2 "Ask CAS" for self-help around the clock  
NEWS 3 FEB 27 New STN AnaVist pricing effective March 1, 2006  
NEWS 4 MAY 10 CA/Capplus enhanced with 1900-1906 U.S. patent records  
NEWS 5 MAY 11 KOREAPAT updates resume  
NEWS 6 MAY 19 Derwent World Patents Index to be reloaded and enhanced  
NEWS 7 MAY 30 IPC 8 Rolled-up Core codes added to CA/Capplus and  
USPATFULL/USPAT2  
NEWS 8 MAY 30 The F-Term thesaurus is now available in CA/Capplus  
NEWS 9 JUN 02 The first reclassification of IPC codes now complete in  
INPADOC  
NEWS 10 JUN 26 TULSA/TULSA2 reloaded and enhanced with new search and  
and display fields  
NEWS 11 JUN 28 Price changes in full-text patent databases EPFULL and PCTFULL  
NEWS 12 JUL 11 CHEMSAFE reloaded and enhanced  
NEWS 13 JUL 14 FSTA enhanced with Japanese patents  
NEWS 14 JUL 19 Coverage of Research Disclosure reinstated in DWPI  
NEWS 15 AUG 09 INSPEC enhanced with 1898-1968 archive  
NEWS 16 AUG 28 ADISCTI Reloaded and Enhanced  
NEWS 17 AUG 30 CA(SM)/Capplus(SM) Austrian patent law changes  
NEWS 18 SEP 11 CA/Capplus enhanced with more pre-1907 records  
NEWS 19 SEP 21 CA/Capplus fields enhanced with simultaneous left and right  
truncation  
NEWS 20 SEP 25 CA(SM)/Capplus(SM) display of CA Lexicon enhanced  
NEWS 21 SEP 25 CAS REGISTRY(SM) no longer includes Concord 3D coordinates  
NEWS 22 SEP 25 CAS REGISTRY(SM) updated with amino acid codes for pyrrolysine  
NEWS 23 SEP 28 CEABA-VTB classification code fields reloaded with new  
classification scheme  
  
NEWS EXPRESS JUNE 30 CURRENT WINDOWS VERSION IS V8.01b, CURRENT  
MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP),  
AND CURRENT DISCOVER FILE IS DATED 26 JUNE 2006.  
  
NEWS HOURS STN Operating Hours Plus Help Desk Availability  
NEWS LOGIN Welcome Banner and News Items  
NEWS IPC8 For general information regarding STN implementation of IPC 8  
NEWS X25 X.25 communication option no longer available

Enter NEWS followed by the item number or name to see news on that  
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\* \* \* \* \* STN Columbus \* \* \* \* \*

FILE 'HOME' ENTERED AT 13:25:58 ON 11 OCT 2006

=>

Uploading

THIS COMMAND NOT AVAILABLE IN THE CURRENT FILE

Do you want to switch to the Registry File?

Choice (Y/n):

Switching to the Registry File...

Some commands only work in certain files. For example, the EXPAND command can only be used to look at the index in a file which has an index. Enter "HELP COMMANDS" at an arrow prompt (=>) for a list of commands which can be used in this file.

=> FILE REGISTRY

COST IN U.S. DOLLARS	SINCE FILE ENTRY	TOTAL SESSION
FULL ESTIMATED COST	0.21	0.21

FILE 'REGISTRY' ENTERED AT 13:26:05 ON 11 OCT 2006

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STRUCTURE FILE UPDATES: 10 OCT 2006 HIGHEST RN 910095-75-9

DICTIONARY FILE UPDATES: 10 OCT 2006 HIGHEST RN 910095-75-9

New CAS Information Use Policies, enter HELP USAGETERMS for details.

TSCA INFORMATION NOW CURRENT THROUGH June 30, 2006

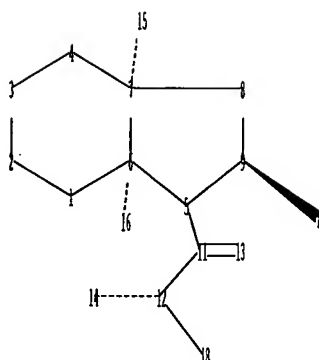
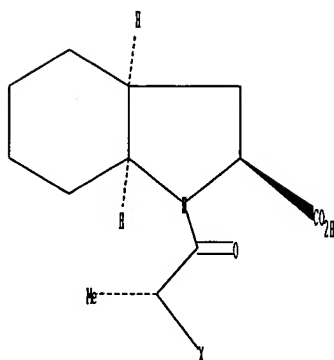
Please note that search-term pricing does apply when conducting SmartSELECT searches.

REGISTRY includes numerically searchable data for experimental and predicted properties as well as tags indicating availability of experimental property data in the original document. For information on property searching in REGISTRY, refer to:

<http://www.cas.org/ONLINE/UG/regprops.html>

=>

Uploading C:\Program Files\Stnexp\Queries\10566562a1.str



chain nodes :  
 10 11 12 13 14 15 16 18  
 ring nodes :  
 1 2 3 4 5 6 7 8 9  
 chain bonds :  
 5-11 6-16 7-15 9-10 11-12 11-13 12-14 12-18  
 ring bonds :  
 1-2 1-6 2-3 3-4 4-7 5-6 5-9 6-7 7-8 8-9  
 exact/norm bonds :  
 5-6 5-9 5-11 6-16 7-15 11-13 12-14  
 exact bonds :  
 1-2 1-6 2-3 3-4 4-7 6-7 7-8 8-9 9-10 11-12 12-18  
 isolated ring systems :  
 containing 1 :

G1:X

Match level :  
 1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:CLASS  
 11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 16:CLASS 18:CLASS

Stereo Bonds:

10-9 (Single Wedge).

Stereo Chiral Centers:

9 (Parity=Don't Care)

Stereo RSS Sets:

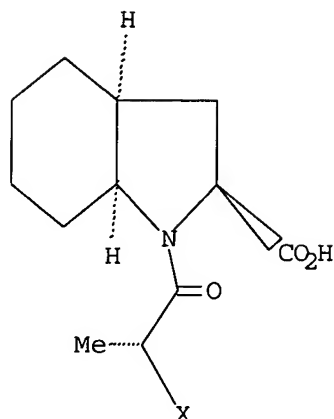
Type=Relative (Default). 1 Nodes= 9

L1 STRUCTURE UPLOADED

=> d 11

L1 HAS NO ANSWERS

L1 STR



G1 X

Structure attributes must be viewed using STN Express query preparation.

=> s l1

SAMPLE SEARCH INITIATED 13:26:22 FILE 'REGISTRY'  
SAMPLE SCREEN SEARCH COMPLETED - 32 TO ITERATE

100.0% PROCESSED 32 ITERATIONS 0 ANSWERS  
SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE \*\*COMPLETE\*\*  
BATCH \*\*COMPLETE\*\*

PROJECTED ITERATIONS: 301 TO 979  
PROJECTED ANSWERS: 0 TO 0

L2 0 SEA SSS SAM L1

=> s l1 sss full

FULL SEARCH INITIATED 13:26:28 FILE 'REGISTRY'  
FULL SCREEN SEARCH COMPLETED - 586 TO ITERATE

100.0% PROCESSED 586 ITERATIONS  
SEARCH TIME: 00.00.01

2 ANSWERS

L3 2 SEA SSS FUL L1

=> FIL HCAPLUS

COST IN U.S. DOLLARS

SINCE FILE	TOTAL
ENTRY	SESSION
166.94	167.15

FULL ESTIMATED COST

FILE 'HCAPLUS' ENTERED AT 13:26:36 ON 11 OCT 2006  
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FILE COVERS 1907 - 11 Oct 2006 VOL 145 ISS 16  
FILE LAST UPDATED: 10 Oct 2006 (20061010/ED)

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This file contains CAS Registry Numbers for easy and accurate substance identification.

=> s l3

L4 1 L3

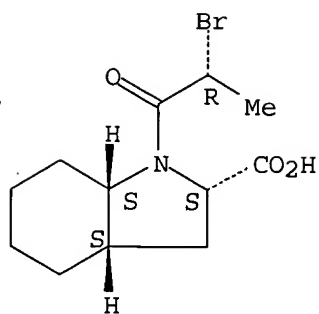
=> d l4 ibib abs hitstr tot

L4 ANSWER 1 OF 1 HCAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 1987:67669 HCAPLUS  
DOCUMENT NUMBER: 106:67669  
TITLE: Indolapril  
INVENTOR(S): Linan Castellet, Isidro; Oliver Mir, Monica  
PATENT ASSIGNEE(S): Farmhispania S. A., Spain; Bioiberica S. A.  
SOURCE: Span., 13 pp.  
CODEN: SPXXAD  
DOCUMENT TYPE: Patent  
LANGUAGE: Spanish  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
ES 537841	A1	19860116	ES 1984-537841	19841121
PRIORITY APPLN. INFO.:			ES 1984-537841	19841121
AB The title compound, useful as an antihypertensive (no data), was prepared An indole-2-carboxylic acid derivative was N-acylated by MeCHBrCOBr and NaHCO <sub>3</sub> and the product was treated with (S)-PhCH <sub>2</sub> CH <sub>2</sub> CH(NH <sub>2</sub> )CO <sub>2</sub> Et and Et <sub>3</sub> N to give Indolapril.				
IT 106534-64-9P				
RL: SPN (Synthetic preparation); PREP (Preparation) (preparation of)				
RN 106534-64-9 HCAPLUS				
CN 1H-Indole-2-carboxylic acid, 1-(2-bromo-1-oxopropyl)octahydro-, [2S-[1(S*),2α,3aβ,7aβ]]- (9CI) (CA INDEX NAME)				

Absolute stereochemistry.



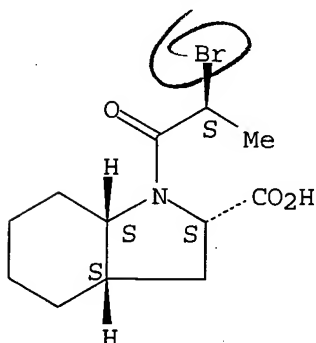
IT 106534-65-0P

RL: SPN (Synthetic preparation); PREP (Preparation)  
 (preparation of, for alkylation of aminobutyric acid derivative)

RN 106534-65-0 HCAPLUS

CN 1H-Indole-2-carboxylic acid, 1-(2-bromo-1-oxopropyl)octahydro-,  
 [2S-[1(R\*),2α,3αβ,7αβ]]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



=&gt; FIL REGISTRY

COST IN U.S. DOLLARS

SINCE FILE	TOTAL
ENTRY	SESSION
15.23	182.38

FULL ESTIMATED COST

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)

SINCE FILE	TOTAL
ENTRY	SESSION
-0.75	-0.75

CA SUBSCRIBER PRICE

FILE 'REGISTRY' ENTERED AT 13:29:05 ON 11 OCT 2006

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DICTIONARY FILE UPDATES: 10 OCT 2006 HIGHEST RN 910095-75-9

New CAS Information Use Policies, enter HELP USAGETERMS for details.

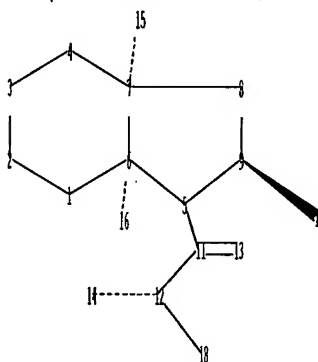
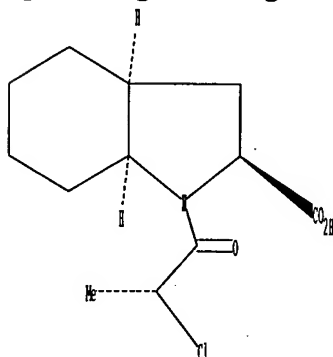
TSCA INFORMATION NOW CURRENT THROUGH June 30, 2006

Please note that search-term pricing does apply when conducting SmartSELECT searches.

REGISTRY includes numerically searchable data for experimental and predicted properties as well as tags indicating availability of experimental property data in the original document. For information on property searching in REGISTRY, refer to:

<http://www.cas.org/ONLINE/UG/regprops.html>

=>  
Uploading C:\Program Files\Stnexp\Queries\10566562a2.str



chain nodes :  
10 11 12 13 14 15 16 18  
ring nodes :  
1 2 3 4 5 6 7 8 9  
chain bonds :  
5-11 6-16 7-15 9-10 11-12 11-13 12-14 12-18  
ring bonds :  
1-2 1-6 2-3 3-4 4-7 5-6 5-9 6-7 7-8 8-9  
exact/norm bonds :  
5-6 5-9 5-11 6-16 7-15 11-13 12-14  
exact bonds :  
1-2 1-6 2-3 3-4 4-7 6-7 7-8 8-9 9-10 11-12 12-18  
isolated ring systems :  
containing 1 :

G1:X

Match level :  
1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:CLASS  
11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 16:CLASS 18:CLASS

Stereo Bonds:

10-9 (Single Wedge).

Stereo Chiral Centers:

9. (Parity=Don't Care)

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Stereo RSS Sets:

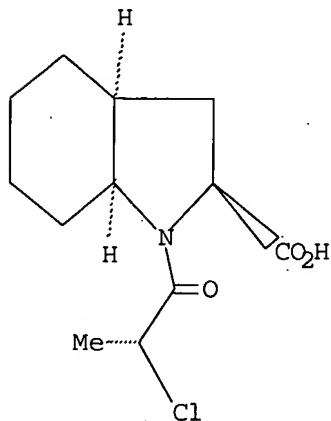
Type=Relative (Default). 1 Nodes= 9

L5 STRUCTURE UPLOADED

=> d 15

L5 HAS NO ANSWERS

L5 STR



G1 X

Structure attributes must be viewed using STN Express query preparation.

=> s 15

SAMPLE SEARCH INITIATED 13:29:24 FILE 'REGISTRY'

SAMPLE SCREEN SEARCH COMPLETED - 2 TO ITERATE

100.0% PROCESSED 2 ITERATIONS

0 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE \*\*COMPLETE\*\*

BATCH \*\*COMPLETE\*\*

PROJECTED ITERATIONS: 2 TO 124

PROJECTED ANSWERS: 0 TO 0

L6 0 SEA SSS SAM L5

=> s 15 sss full

FULL SEARCH INITIATED 13:29:30 FILE 'REGISTRY'

FULL SCREEN SEARCH COMPLETED - 45 TO ITERATE

100.0% PROCESSED 45 ITERATIONS

0 ANSWERS

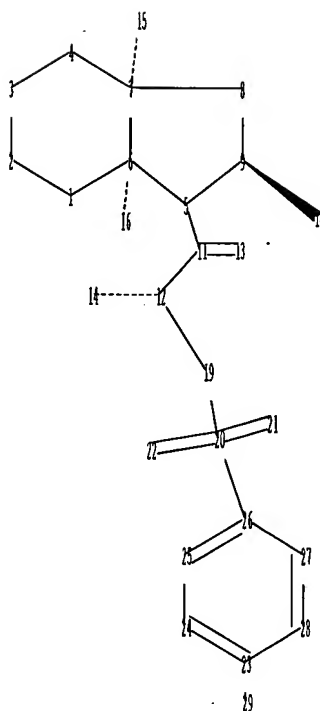
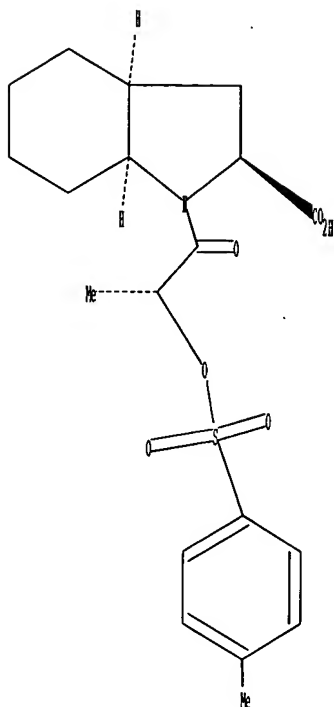
SEARCH TIME: 00.00.01

L7 0 SEA SSS FUL L5

=>

Uploading C:\Program Files\Stnexp\Queries\10566561a3.str





chain nodes :

10 11 12 13 14 15 16 19 20 21 22 29

ring nodes :

1 2 3 4 5 6 7 8 9 23 24 25 26 27 28

chain bonds :

5-11 6-16 7-15 9-10 11-12 11-13 12-14 12-19 19-20 20-21 20-22 20-26  
23-29

ring bonds :

1-2 1-6 2-3 3-4 4-7 5-6 5-9 6-7 7-8 8-9 23-24 23-28 24-25 25-26 26-27  
27-28

exact/norm bonds :

5-6 5-9 5-11 6-16 7-15 11-13 12-14 12-19 19-20 20-21 20-22 20-26

exact bonds :

1-2 1-6 2-3 3-4 4-7 6-7 7-8 8-9 9-10 11-12 23-29

normalized bonds :

23-24 23-28 24-25 25-26 26-27 27-28

isolated ring systems :

containing 1 : 23 :

G1:X

Match level :

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:CLASS  
11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 16:CLASS 19:CLASS 20:CLASS  
21:CLASS 22:CLASS 23:Atom 24:Atom 25:Atom 26:Atom 27:Atom 28:Atom 29:CLASS

Stereo Bonds:

10-9 (Single Wedge).

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Stereo Chiral Centers:

9 (Parity=Don't Care)

Stereo RSS Sets:

Type=Relative (Default). 1 Nodes= 9

L8 STRUCTURE UPLOADED

=> d l8

L8 HAS NO ANSWERS

L8 STR

\* STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT \*

Structure attributes must be viewed using STN Express query preparation.

=> s l8

SAMPLE SEARCH INITIATED 13:33:31 FILE 'REGISTRY'

SAMPLE SCREEN SEARCH COMPLETED - 1 TO ITERATE

100.0% PROCESSED 1 ITERATIONS

0 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE \*\*COMPLETE\*\*

BATCH \*\*COMPLETE\*\*

PROJECTED ITERATIONS: 1 TO 80

PROJECTED ANSWERS: 0 TO 0

L9 0 SEA SSS SAM L8

=> s l8 sss full

FULL SEARCH INITIATED 13:33:37 FILE 'REGISTRY'

FULL SCREEN SEARCH COMPLETED - 12 TO ITERATE

100.0% PROCESSED 12 ITERATIONS

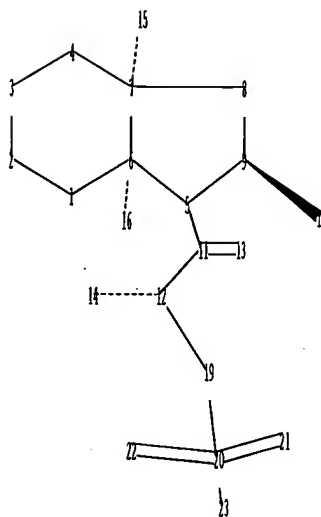
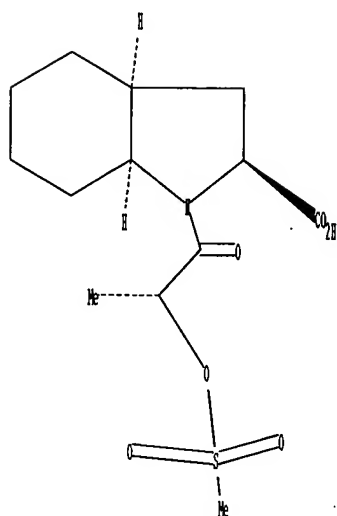
0 ANSWERS

SEARCH TIME: 00.00.01

L10 0 SEA SSS FUL L8

=>

Uploading C:\Program Files\Stnexp\Queries\10566562a4.str



chain nodes :

10 11 12 13 14 15 16 19 20 21 22 23

ring nodes :

1 2 3 4 5 6 7 8 9

chain bonds :

5-11 6-16 7-15 9-10 11-12 11-13 12-14 12-19 19-20 20-21 20-22 20-23

ring bonds :

1-2 1-6 2-3 3-4 4-7 5-6 5-9 6-7 7-8 8-9

exact/norm bonds :

5-6 5-9 5-11 6-16 7-15 11-13 12-14 12-19 19-20 20-21 20-22

exact bonds :

1-2 1-6 2-3 3-4 4-7 6-7 7-8 8-9 9-10 11-12 20-23

isolated ring systems :

containing 1 :

G1:X

Match level :

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:CLASS  
11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 16:CLASS 19:CLASS 20:CLASS  
21:CLASS 22:CLASS 23:CLASS

Stereo Bonds:

10-9 (Single Wedge).

Stereo Chiral Centers:

9 (Parity=Don't Care)

Stereo RSS Sets:

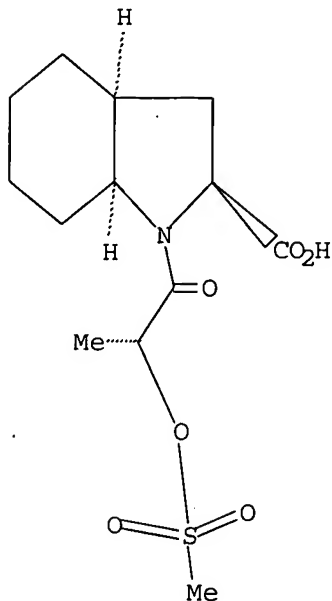
Type=Relative (Default). 1 Nodes= 9

L11 STRUCTURE UPLOADED

=> d l11

10/11/2006 10566562a1.trn

L11 HAS NO ANSWERS  
L11 STR



G1 X

Structure attributes must be viewed using STN Express query preparation.

=> s l11

SAMPLE SEARCH INITIATED 13:36:57 FILE 'REGISTRY'  
SAMPLE SCREEN SEARCH COMPLETED - 1 TO ITERATE

100.0% PROCESSED 1 ITERATIONS 0 ANSWERS  
SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE \*\*COMPLETE\*\*  
BATCH \*\*COMPLETE\*\*  
PROJECTED ITERATIONS: 1 TO 80  
PROJECTED ANSWERS: 0 TO 0

L12 0 SEA SSS SAM L11

=> s l11 sss full

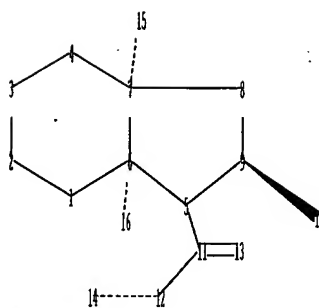
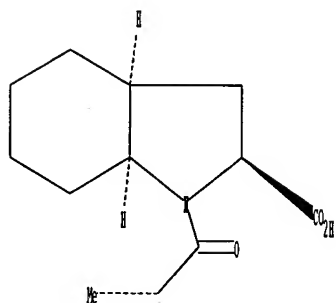
FULL SEARCH INITIATED 13:37:03 FILE 'REGISTRY'  
FULL SCREEN SEARCH COMPLETED - 12 TO ITERATE

100.0% PROCESSED 12 ITERATIONS 0 ANSWERS  
SEARCH TIME: 00.00.01

L13 0 SEA SSS FUL L11

=>

Uploading C:\Program Files\Stnexp\Queries\10566562a5.str



chain nodes :  
 10 11 12 13 14 15 16  
 ring nodes :  
 1 2 3 4 5 6 7 8 9  
 chain bonds :  
 5-11 6-16 7-15 9-10 11-12 11-13 12-14  
 ring bonds :  
 1-2 1-6 2-3 3-4 4-7 5-6 5-9 6-7 7-8 8-9  
 exact/norm bonds :  
 5-6 5-9 5-11 6-16 7-15 11-13 12-14  
 exact bonds :  
 1-2 1-6 2-3 3-4 4-7 6-7 7-8 8-9 9-10 11-12  
 isolated ring systems :  
 containing 1 :

G1:X

Match level :  
 1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:CLASS  
 11:CLASS 12:CLASS 13:CLASS 14:CLASS 15:CLASS 16:CLASS

Stereo Bonds:

10-9 (Single Wedge).

Stereo Chiral Centers:

9 (Parity=Don't Care)

Stereo RSS Sets:

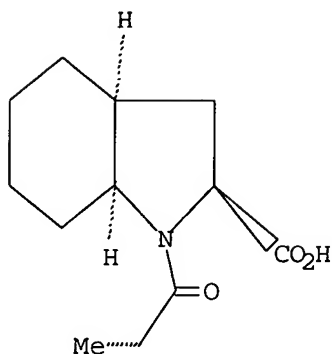
Type=Relative (Default). 1 Nodes= 9

L14 STRUCTURE UPLOADED

=> d l14

L14 HAS NO ANSWERS

L14 STR



G1 X

Structure attributes must be viewed using STN Express query preparation.

=> s l14

SAMPLE SEARCH INITIATED 13:40:03 FILE 'REGISTRY'  
SAMPLE SCREEN SEARCH COMPLETED - 143 TO ITERATE

100.0% PROCESSED 143 ITERATIONS 12 ANSWERS  
SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE \*\*COMPLETE\*\*  
BATCH \*\*COMPLETE\*\*  
PROJECTED ITERATIONS: 2143 TO 3577  
PROJECTED ANSWERS: 33 TO 447

L15 12 SEA SSS SAM L14

=> s l14 sss full

FULL SEARCH INITIATED 13:40:10 FILE 'REGISTRY'  
FULL SCREEN SEARCH COMPLETED - 3181 TO ITERATE

100.0% PROCESSED 3181 ITERATIONS 243 ANSWERS  
SEARCH TIME: 00.00.01

L16 243 SEA SSS FUL L14

=> FIL HCAPLUS

COST IN U.S. DOLLARS	SINCE FILE ENTRY	TOTAL SESSION
FULL ESTIMATED COST	674.80	857.18
DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)	SINCE FILE ENTRY	TOTAL SESSION
CA SUBSCRIBER PRICE	0.00	-0.75

FILE 'HCAPLUS' ENTERED AT 13:40:51 ON 11 OCT 2006  
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FILE COVERS 1907 - 11 Oct 2006 VOL 145 ISS 16  
FILE LAST UPDATED: 10 Oct 2006 (20061010/ED)

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This file contains CAS Registry Numbers for easy and accurate substance identification.

=> d his

(FILE 'HOME' ENTERED AT 13:25:58 ON 11 OCT 2006)

FILE 'REGISTRY' ENTERED AT 13:26:05 ON 11 OCT 2006

L1 STRUCTURE UPLOADED  
L2 0 S L1  
L3 2 S L1 SSS FULL

FILE 'HCAPLUS' ENTERED AT 13:26:36 ON 11 OCT 2006

L4 1 S L3

FILE 'REGISTRY' ENTERED AT 13:29:05 ON 11 OCT 2006

L5 STRUCTURE UPLOADED  
L6 0 S L5  
L7 0 S L5 SSS FULL  
L8 STRUCTURE UPLOADED  
L9 0 S L8  
L10 0 S L8 SSS FULL  
L11 STRUCTURE UPLOADED  
L12 0 S L11  
L13 0 S L11 SSS FULL  
L14 STRUCTURE UPLOADED  
L15 12 S L14  
L16 243 S L14 SSS FULL

FILE 'HCAPLUS' ENTERED AT 13:40:51 ON 11 OCT 2006

=> s 116

L17 1521 L16

=> s 117 and perindopril

1113 PERINDOPRIL

L18 983 L17 AND PERINDOPRIL

=> s 118 and process

2320069 PROCESS

1574509 PROCESSES

3462426 PROCESS

(PROCESS OR PROCESSES)

L19 68 L18 AND PROCESS

=> s l19 and synthesis

1272425 SYNTHESIS

3 SYNTHESISES

67891 SYNTHESES

1 SYNTHESISES

1310499 SYNTHESIS

(SYNTHESIS OR SYNTHESISES OR SYNTHESES OR SYNTHESISES)

L20

12 L19 AND SYNTHESIS

=> d l20 ibib abs hitstr tot

L20 ANSWER 1 OF 12 HCAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2006:796623 HCAPLUS

DOCUMENT NUMBER: 145:230528

TITLE: Process for making highly pure  
perindopril erbumine

INVENTOR(S): Kumar, Ashok; Soudagar, Satish Rajanikant; Mathur,  
Arpana; Shah, Chirag Hasmukh; Gunjal, Sanjay Tukaram;  
Metil, Dattatray Shamrao; Kelkar, Rahul Suresh;  
Thakare, Devendra Digambar; Kumar, Bindu Manoj; Nair,  
Raji

PATENT ASSIGNEE(S): USA

SOURCE: U.S. Pat. Appl. Publ., 6pp.

CODEN: USXXCO

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 2006178422	A1	20060810	US 2005-140226	20050527
PRIORITY APPLN. INFO.:			IN 2004-MU566	A 20040531
OTHER SOURCE(S):	CASREACT 145:230528			

AB A process for the synthesis and isolation of  
(2S,3aS,7aS)-1-[(2S)-2-[[[(1S)-1-(ethoxycarbonyl)butyl]amino]-1-oxopropyl]octahydro-1H-indole-2-carboxylic acid and its tert-butylamine salt, comprises the amidation of (2S,3aS,7aS)-octahydroindole-2-carboxylic acid benzyl ester and N-[(S)1-carboxybutyl]-(S)-alanine Et ester in nonreactive solvents in turn avoiding the formation of the impurity N-acetyl (2S,3aS,7aS)-octahydroindole-2-carboxylic acid benzyl ester. The de-protection of benzyl ester group is optimized by catalytic hydrogenolysis and then isolation of the product from an aqueous layer by extraction using an organic solvent, which eliminates the need for lyophilization.

This yields perindopril erbumine free of contaminants derivable from dicyclohexylcarbodiimide (e.g., dicyclohexylurea) and impurities originated by the use of Et acetate.

IT 82834-16-0P, Perindopril

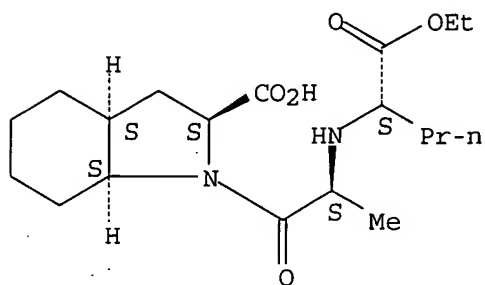
RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)  
(in a process for making highly pure perindopril erbumine)

RN 82834-16-0 HCAPLUS

CN 1H-Indole-2-carboxylic acid, 1-[(2S)-2-[[[(1S)-1-(ethoxycarbonyl)butyl]amino]-1-oxopropyl]octahydro-, (2S,3aS,7aS)- (9CI)  
(CA INDEX NAME)

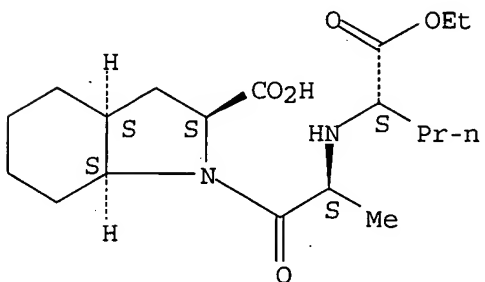


Absolute stereochemistry. Rotation (-).

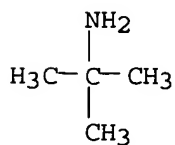


IT 107133-36-8P, Perindopril erbumine  
 RL: SPN (Synthetic preparation); PREP (Preparation)  
 (process for making highly pure perindopril  
 erbumine)  
 RN 107133-36-8 HCAPLUS  
 CN 1H-Indole-2-carboxylic acid, 1-[(2S)-2-[[[(1S)-1-(ethoxycarbonyl)butyl]amino]-1-oxopropyl]octahydro-, (2S,3aS,7aS)-, compd.  
 with 2-methyl-2-propanamine (1:1) (9CI) (CA INDEX NAME)  
 CM 1  
 CRN 82834-16-0  
 CMF C19 H32 N2 O5

Absolute stereochemistry. Rotation (-).



CM 2  
 CRN 75-64-9  
 CMF C4 H11 N



L20 ANSWER 2 OF 12 HCAPLUS COPYRIGHT 2006 ACS on STN  
 ACCESSION NUMBER: 2006:680403 HCAPLUS

DOCUMENT NUMBER: 145:124844  
 TITLE: Process for the synthesis of  
 (2S,3aS,7aS)-1-(S)-alanyloctahydro-1H-indole-2-  
 carboxylic acid derivatives and use in the  
 synthesis of perindopril  
 INVENTOR(S): Kumar, Ashok; Soudagar, Satish Rajanikant; Mathur,  
 Arpana; Gunjal, Sanjay Tukaram; Panda, Nalinakshya  
 Balaram; Jadhav, Dilip Uttam  
 PATENT ASSIGNEE(S): IPCA Laboratories Limited, India  
 SOURCE: Eur. Pat. Appl., 16 pp.  
 CODEN: EPXXDW  
 DOCUMENT TYPE: Patent  
 LANGUAGE: English  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 1679072	A1	20060712	EP 2005-113099	20051230
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, EI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK, BA, HR, IS, YU				

PRIORITY APPLN. INFO.: IN 2005-MU17 A 20050106  
 OTHER SOURCE(S): CASREACT 145:124844

AB The invention relates perindopril [(2S,3aS,7aS)-1-[(2S)-2-[(S)-1-(ethoxycarbonyl)butylamino]propionyl]octahydro-1H-indole-2-carboxylic acid] aralkyl ester salts used in the synthesis of perindopril. Thus, (2S,3aS,7aS)-octahydro-1H-indole-2-carboxylic acid was treated with N-[(S)-1-(ethoxycarbonyl)butyl]-L-alanine in CH<sub>2</sub>Cl<sub>2</sub> in the presence of Et<sub>3</sub>N, 1-hydroxybenzotriazole, and dicyclohexylcarbodiimide to afford 99% perindopril benzyl ester. Conversion of the latter into the oxalate salt, followed by hydrogenolysis over 5% Pd/C and reaction with tert-butylamine yielded perindopril erbumine.

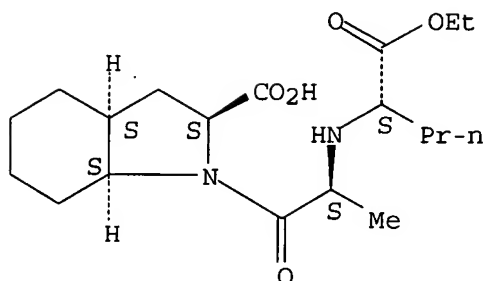
IT 107133-36-8P, Perindopril erbumine  
 RL: IMF (Industrial manufacture); SPN (Synthetic preparation); PREP (Preparation)  
 (process for synthesis of  
 alanyloctahydroindolecarboxylic acid derivs. in synthesis of  
 perindopril)

RN 107133-36-8 HCAPLUS  
 CN 1H-Indole-2-carboxylic acid, 1-[(2S)-2-[[1S)-1-(ethoxycarbonyl)butyl]amino]-1-oxopropyl]octahydro-, (2S,3aS,7aS)-, compd. with 2-methyl-2-propanamine (1:1) (9CI) (CA INDEX NAME)

CM 1

CRN 82834-16-0  
 CMF C19 H32 N2 O5

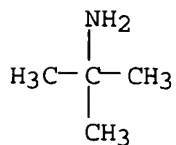
Absolute stereochemistry. Rotation (-).



CM 2

CRN 75-64-9

CMF C4 H11 N



IT 897922-10-0P 897922-12-2P 897922-14-4P

897922-16-6P 897922-21-3P 897922-22-4P

897922-23-5P 897922-25-7P 897922-27-9P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(process for synthesis of

alanyloctahydroindolecarboxylic acid derivs. in synthesis of perindopril)

RN 897922-10-0 HCAPLUS

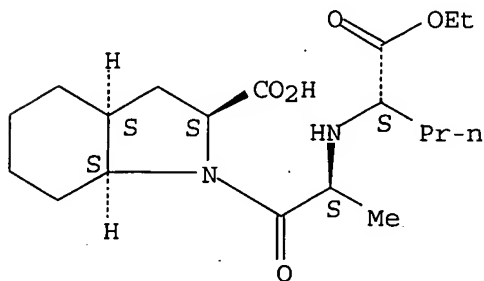
CN 1H-Indole-2-carboxylic acid, 1-[(2S)-2-[[[(1S)-1-(ethoxycarbonyl)butyl]amino]-1-oxopropyl]octahydro-, (2S,3aS,7aS)-, ethanedioate (9CI) (CA INDEX NAME)

CM 1

CRN 82834-16-0

CMF C19 H32 N2 O5

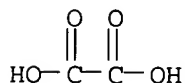
Absolute stereochemistry. Rotation (-).



CM 2

CRN 144-62-7

CMF C2 H2 O4



RN 897922-12-2 HCAPLUS

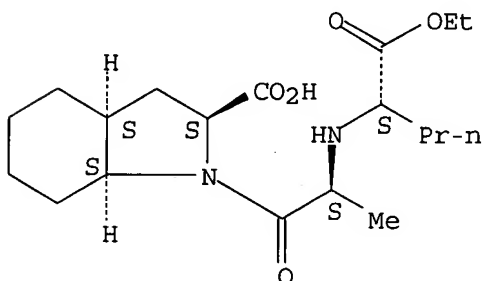
CN Butanedioic acid, 2,3-bis[(4-methylbenzoyl)oxy]-, (2R,3R)-, compd. with (2S,3aS,7aS)-1-[(2S)-2-[[[(1S)-1-(ethoxycarbonyl)butyl]amino]-1-oxopropyl]octahydro-1H-indole-2-carboxylic acid (9CI) (CA INDEX NAME)

CM 1

CRN 82834-16-0

CMF C19 H32 N2 O5

Absolute stereochemistry. Rotation (-).

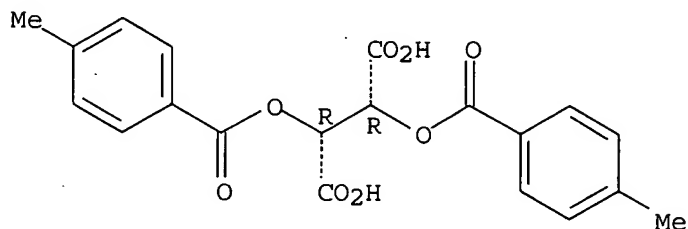


CM 2

CRN 32634-66-5

CMF C20 H18 O8

Absolute stereochemistry. Rotation (-).



RN 897922-14-4 HCAPLUS

CN Butanedioic acid, 2,3-dihydroxy- (2R,3R)-, compd. with (2S,3aS,7aS)-1-[(2S)-2-[[[(1S)-1-(ethoxycarbonyl)butyl]amino]-1-oxopropyl]octahydro-1H-indole-2-carboxylic acid (9CI) (CA INDEX NAME)

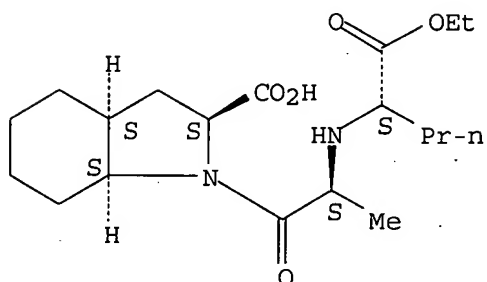
10/11/2006 10566562a1.trn

CM 1

CRN 82834-16-0

CMF C19 H32 N2 O5

Absolute stereochemistry. Rotation (-).

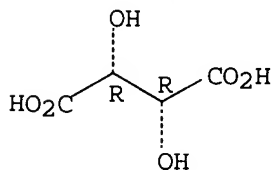


CM 2

CRN 87-69-4

CMF C4 H6 O6

Absolute stereochemistry.



RN 897922-16-6 HCAPLUS

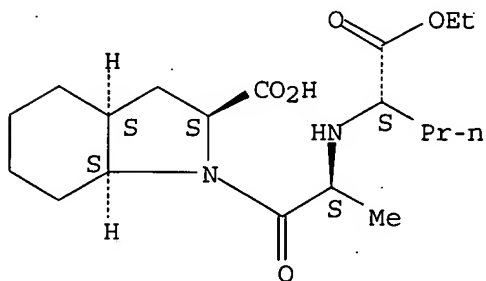
CN 1H-Indole-2-carboxylic acid, 1-[(2S)-2-[[[(1S)-1-(ethoxycarbonyl)butyl]amino]-1-oxopropyl]octahydro-, (2S,3aS,7aS)-, phosphate (9CI) (CA INDEX NAME)

CM 1

CRN 82834-16-0

CMF C19 H32 N2 O5

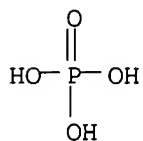
Absolute stereochemistry. Rotation (-).



CM 2

CRN 7664-38-2

CMF H3 O4 P



RN 897922-21-3 HCAPLUS

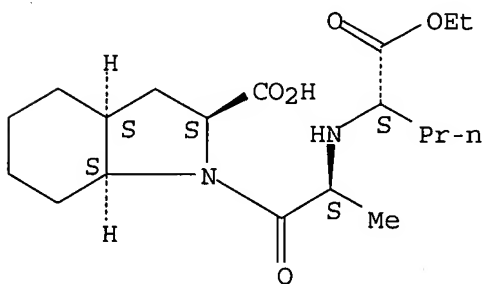
CN 1,2-Benzenedicarboxylic acid, compd. with (2S,3aS,7aS)-1-[(2S)-2-[[[(1S)-1-(ethoxycarbonyl)butyl]amino]-1-oxopropyl]octahydro-1H-indole-2-carboxylic acid (9CI) (CA INDEX NAME)

CM 1

CRN 82834-16-0

CMF C19 H32 N2 O5

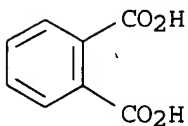
Absolute stereochemistry. Rotation (-).



CM 2

CRN 88-99-3

CMF C8 H6 O4



RN 897922-22-4 HCAPLUS

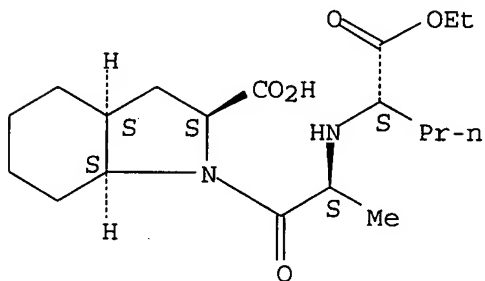
CN 1H-Indole-2-carboxylic acid, 1-[(2S)-2-[[[(1S)-1-(ethoxycarbonyl)butyl]amino]-1-oxopropyl]octahydro-, (2S,3aS,7aS)-, (1S,4R)-7,7-dimethyl-2-oxobicyclo[2.2.1]heptane-1-methanesulfonate (9CI) (CA INDEX NAME)

CM 1

10/11/2006 10566562a1.trn

CRN 82834-16-0  
CMF C19 H32 N2 O5

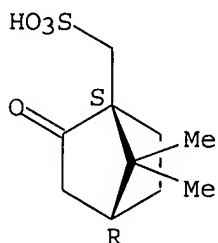
Absolute stereochemistry. Rotation (-).



CM 2

CRN 3144-16-9  
CMF C10 H16 O4 S

Absolute stereochemistry. Rotation (+).



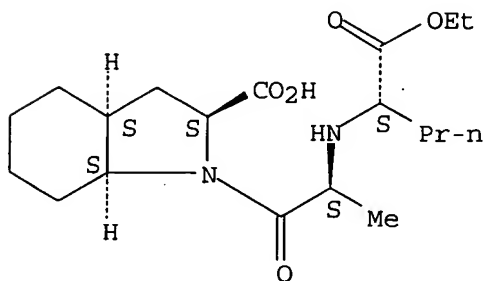
RN 897922-23-5 HCAPLUS

CN 1H-Indole-2-carboxylic acid, 1-[[ (2S)-2-[[ (1S)-1-(ethoxycarbonyl)butyl]amino]-1-oxopropyl]octahydro-, (2S,3aS,7aS)-, 2-hydroxy-1,2,3-propanetricarboxylate (9CI) (CA INDEX NAME)

CM 1

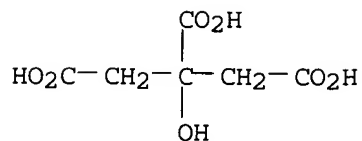
CRN 82834-16-0  
CMF C19 H32 N2 O5

Absolute stereochemistry. Rotation (-).



CM 2

CRN 77-92-9  
CMF C6 H8 O7

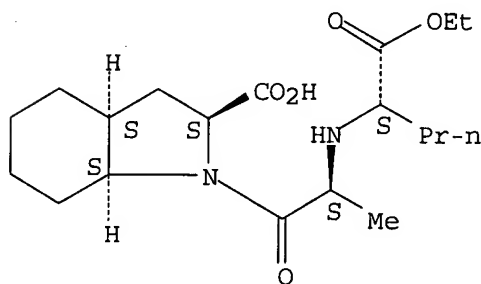


RN 897922-25-7 HCAPLUS  
CN Butanedioic acid, 2,3-bis[(4-methylbenzoyl)oxy]-, (2S,3S)-, compd. with (2S,3aS,7aS)-1-[(2S)-2-[[[(1S)-1-(ethoxycarbonyl)butyl]amino]-1-oxopropyl]octahydro-1H-indole-2-carboxylic acid (9CI) (CA INDEX NAME)

CM 1

CRN 82834-16-0  
CMF C19 H32 N2 O5

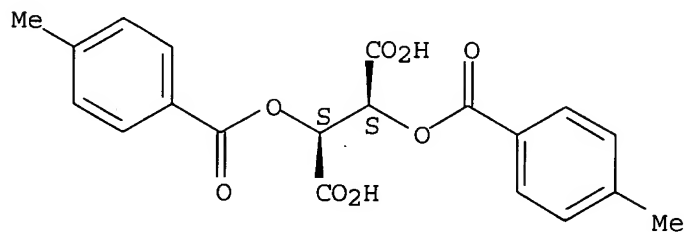
Absolute stereochemistry. Rotation (-).



CM 2

CRN 32634-68-7  
CMF C20 H18 O8

Absolute stereochemistry. Rotation (+).



RN 897922-27-9 HCAPLUS  
CN Butanedioic acid, 2,3-bis(benzoyloxy)-, (2R,3R)-rel-, compd. with (2S,3aS,7aS)-1-[(2S)-2-[[[(1S)-1-(ethoxycarbonyl)butyl]amino]-1-



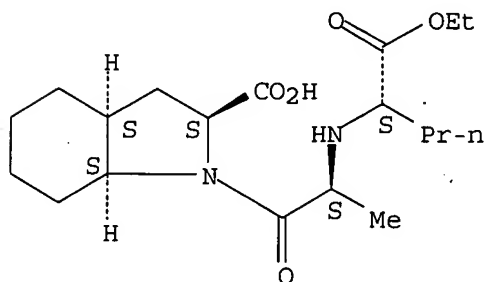
oxopropyl]octahydro-1H-indole-2-carboxylic acid (9CI) (CA INDEX NAME)

CM 1

CRN 82834-16-0

CMF C19 H32 N2 O5

Absolute stereochemistry. Rotation (-).

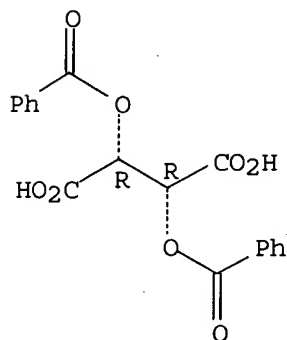


CM 2

CRN 22333-70-6

CMF C18 H14 O8

Relative stereochemistry.



REFERENCE COUNT: 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L20 ANSWER 3 OF 12 HCAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2005:1262577 HCAPLUS

DOCUMENT NUMBER: 144:7098

TITLE: Process for the preparation of perindopril and its salts

INVENTOR(S): Merslavic, Marjo; Smid, Janja; Tomsic, Zdenka

PATENT ASSIGNEE(S): Krka, Tovarna Zdravil D.D. Novo Mesto, Slovenia

SOURCE: PCT Int. Appl., 19 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

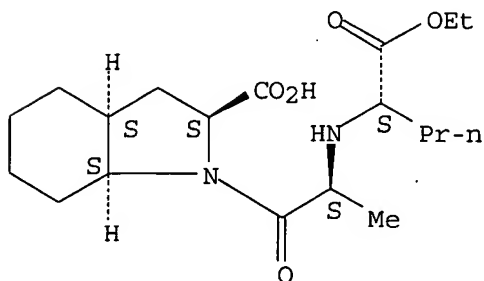
LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2005113500	A1	20051201	WO. 2005-EP5048	20050510
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW				
RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
SI 21800	C	20051231	SI 2004-143	20040514
SI 21852	C	20060228	SI 2004-235	20040805
PRIORITY APPLN. INFO.:			SI 2004-143	A 20040514
			SI 2004-235	A 20040805
OTHER SOURCE(S): CASREACT 144:7098; MARPAT 144:7098				
AB The invention relates to a process for the preparation of the ACE inhibitor perindopril, its pharmaceutically-acceptable salts and intermediates obtained in the process. The process involves conversion of N-[(1S)-1-carbethoxybutyl]-L-alanine to the acid chloride hydrochloride and reaction with (2S,3aS,7aS)-octahydroindole-2-carboxylic acid or a an ester or salt. The examples describe the synthesis of perindopril erbumine by reactions carried out in CH <sub>2</sub> Cl <sub>2</sub> .				
IT 82834-16-0P, Perindopril 107133-36-8P, Perindopril erbumine 869954-04-1P 869954-08-5P 869954-09-6P				
RL: IMF (Industrial manufacture); SPN (Synthetic preparation); PREP (Preparation) (process for preparation of perindopril and its salts)				
RN 82834-16-0 HCAPLUS				
CN 1H-Indole-2-carboxylic acid, 1-[(2S)-2-[[[(1S)-1-(ethoxycarbonyl)butyl]amino]-1-oxopropyl]octahydro-, (2S,3aS,7aS)- (9CI) (CA INDEX NAME)				

Absolute stereochemistry. Rotation (-).



RN 107133-36-8 HCAPLUS  
 CN 1H-Indole-2-carboxylic acid, 1-[(2S)-2-[[[(1S)-1-(ethoxycarbonyl)butyl]amino]-1-oxopropyl]octahydro-, (2S,3aS,7aS)-, compd. with 2-methyl-2-propanamine (1:1) (9CI) (CA INDEX NAME)

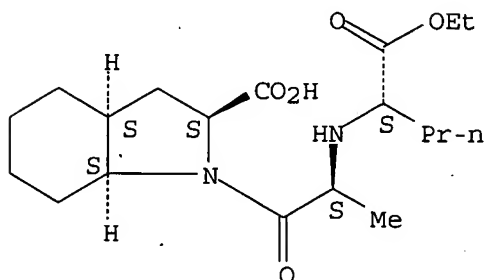
10/11/2006 10566562a1.trn

CM 1

CRN 82834-16-0

CMF C19 H32 N2 O5

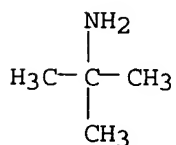
Absolute stereochemistry. Rotation (-).



CM 2

CRN 75-64-9

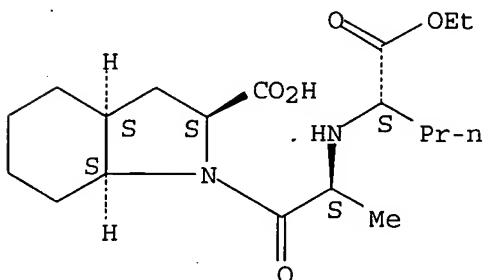
CMF C4 H11 N



RN 869954-04-1 HCAPLUS

CN 1H-Indole-2-carboxylic acid, 1-[(2S)-2-[[[(1S)-1-(ethoxycarbonyl)butyl]amino]-1-oxopropyl]octahydro-, monopotassium salt, (2S,3aS,7aS)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).



● K

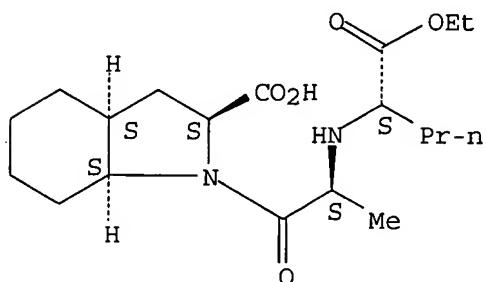
RN 869954-08-5 HCAPLUS

CN 1H-Indole-2-carboxylic acid, 1-[(2S)-2-[[[(1S)-1-(ethoxycarbonyl)butyl]amino]-1-oxopropyl]octahydro-, monolithium salt,

10/11/2006 10566562a1.trn

(2S,3aS,7aS) - (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

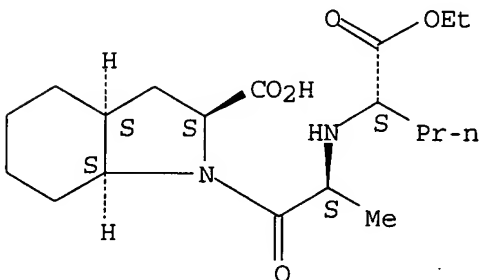


● Li

RN 869954-09-6 HCAPLUS

CN 1H-Indole-2-carboxylic acid, 1-[(2S)-2-[[[(1S)-1-(ethoxycarbonyl)butyl]amino]-1-oxopropyl]octahydro-, monosodium salt, (2S,3aS,7aS) - (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).



● Na

REFERENCE COUNT: 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L20 ANSWER 4 OF 12 HCAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2005:1117891 HCAPLUS

DOCUMENT NUMBER: 143:367597

TITLE: Process for the preparation of perindopril

INVENTOR(S): Kankan, Rajendra Narayanrao; Rao, Dharmaraj Ramachandra

PATENT ASSIGNEE(S): Neopharma Limited, UK

SOURCE: Brit. UK Pat. Appl., 21 pp.

CODEN: BAXXDU

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

## PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
GB 2413128	A1	20051019	GB 2004-8258	20040413
WO 2005100317	A1	20051027	WO 2005-GB1355	20050407

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW

RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG

PRIORITY APPLN. INFO.: GB 2004-8258 A 20040413

OTHER SOURCE(S): MARPAT 143:367597

AB A process for preparing perindopril or a pharmaceutically-acceptable salt comprises coupling a 4-halo-, 4-alkoxy- or 4-nitrobenzyl ester of (2S,3aS,7aS)-2-carboxyoctahydroindole with N-[(S)-1-carbethoxybutyl]-L-alanine (1) in the presence of DCC and HOBT, followed by catalytic hydrolysis. The starting ester was obtained from (S)-indoline-2-carboxylic acid by hydrogenation-esterification and 1 was obtained from norvaline Et ester and pyruvic acid under catalytic hydrogenation conditions. The method was applied to the synthesis perindopril erbumine (20.5 g obtained from 24 g 4-chlorobenzyl ester and 21.26 g 1).

IT 82834-16-0P, Perindopril 107133-36-8P,  
Perindopril erbumine

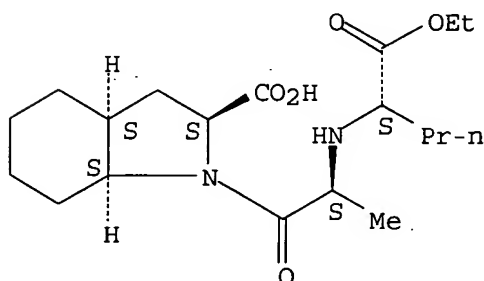
RL: IMF (Industrial manufacture); SPN (Synthetic preparation); PREP (Preparation)

(preparation of perindopril by acylation of octahydroindolecarboxylates with ethoxycarbonylbutylalanine)

RN 82834-16-0 HCAPLUS

CN 1H-Indole-2-carboxylic acid, 1-[(2S)-2-[[[(1S)-1-(ethoxycarbonyl)butyl]amino]-1-oxopropyl]octahydro-, (2S,3aS,7aS)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).



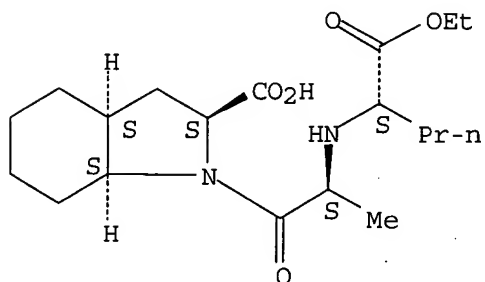
RN 107133-36-8 HCAPLUS

CN 1H-Indole-2-carboxylic acid, 1-[(2S)-2-[[[(1S)-1-(ethoxycarbonyl)butyl]amino]-1-oxopropyl]octahydro-, (2S,3aS,7aS)-, compd. with 2-methyl-2-propanamine (1:1) (9CI) (CA INDEX NAME)

CM 1

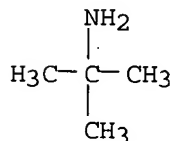
CRN 82834-16-0  
CMF C19 H32 N2 O5

Absolute stereochemistry. Rotation (-).



CM 2

CRN 75-64-9  
CMF C4 H11 N



REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L20 ANSWER 5 OF 12 HCAPLUS COPYRIGHT 2006 ACS on STN  
ACCESSION NUMBER: 2004:1016010 HCAPLUS  
DOCUMENT NUMBER: 141:424441  
TITLE: Process for the preparation of enalapril maleate and related compounds having ACE inhibitory action  
INVENTOR(S): Jenko, Branko  
PATENT ASSIGNEE(S): Lek Pharmaceuticals D.D., Slovenia  
SOURCE: PCT Int. Appl., 18 pp.  
CODEN: PIXXD2  
DOCUMENT TYPE: Patent  
LANGUAGE: English  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2004101515	A1	20041125	WO 2004-SI21	20040507
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,				

NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,  
 TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW  
 RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM,  
 AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK,  
 EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE,  
 SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE,  
 SN, TD, TG

SI 21507 C 20041231 SI 2003-123 20030516  
 EP 1628956 A1 20060301 EP 2004-731808 20040507

R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,  
 IE, SI, FI, RO, CY, TR, BG, CZ, EE, HU, PL, SK

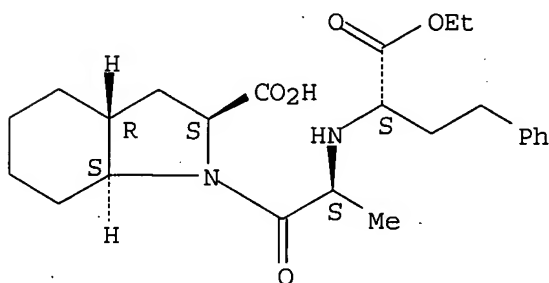
PRIORITY APPLN. INFO.:

SI 2003-123 A 20030516  
 WO 2004-SI21 W 20040507

OTHER SOURCE(S): MARPAT 141:424441

AB The invention relates to a process for the preparation of  
 ACE-inhibitory peptides (S,S)-R1CH2CH2CH(CO2R2)-L-Ala-NR3R4 (R1 is H,  
 alkyl, phenyl; R2 is H, alkyl; NR3R4 is a proline, 2-piperidinecarboxylic  
 or hexahydro-2-azepinecarboxylic acid residue and related aza/thia analogs  
 and their esters or metal salts) in which the carboxy group of  
 (S,S)-R1CH2CH2CH(CO2R2)-L-Ala-OH is activated with a uronium salt in an  
 aprotic solvent prior to coupling with an amino acid HNR3R4. Thus, a  
 mixture of N-[1(S)-(ethoxycarbonyl)-3-phenylpropyl]-L-alanine, L-proline,  
 Et3N and O-(benzotriazol-1-yl)-N,N,N'-tetramethyluronium  
 hexafluorophosphate in acetonitrile-DMF was stirred for 30 min at room  
 temperature to afford enalapril (85.4% yield of maleate).  
 IT 87679-37-6P, Trandolapril  
 RL: IMF (Industrial manufacture); SPN (Synthetic preparation); PREP  
 (Preparation)  
 (preparation of enalapril maleate and related compds. having ACE inhibitory  
 action)  
 RN 87679-37-6 HCAPLUS  
 CN 1H-Indole-2-carboxylic acid, 1-[(2S)-2-[[[(1S)-1-(ethoxycarbonyl)-3-  
 phenylpropyl]amino]-1-oxopropyl]octahydro-, (2S,3aR,7aS)- (9CI) (CA INDEX  
 NAME)

Absolute stereochemistry. Rotation (-).



REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS  
 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L20 ANSWER 6 OF 12 HCAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2004:266899 HCAPLUS

DOCUMENT NUMBER: 140:253919

TITLE: Process for the synthesis of  
 N-[(S)-1-(ethoxycarbonyl)butyl]-L-alanine for use in  
 the synthesis of perindopril  
 INVENTOR(S): Breard, Fabienne; Lecouve, Jean-Pierre

PATENT ASSIGNEE(S): Les Laboratoires Servier, Fr.  
SOURCE: Eur. Pat. Appl., 9 pp.  
CODEN: EPXXDW  
DOCUMENT TYPE: Patent  
LANGUAGE: French  
FAMILY ACC. NUM. COUNT: 1  
PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 1403278	A1	20040331	EP 2003-292404	20030930
EP 1403278	B1	20050608		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK				
AT 297407	E	20050615	AT 2003-292404	20030930
PT 1403278	T	20050930	PT 2003-292404	20030930
ES 2240926	T3	20051016	ES 2003-3292404	20030930
WO 2005033127	A1	20050414	WO 2004-FR2463	20040929
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW				
RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				

PRIORITY APPLN. INFO.: EP 2003-292404 A 20030930

OTHER SOURCE(S): MARPAT 140:253919

AB Perindopril intermediate (S)-EtO<sub>2</sub>CCHPr-L-Ala-OH was prepared by condensation of L-alanine alkyl or benzyl ester with Et glyoxylate or Et chloro(cyclohexyloxy)acetate, followed by allylation with allylzinc bromide, and catalytic hydrogenation.

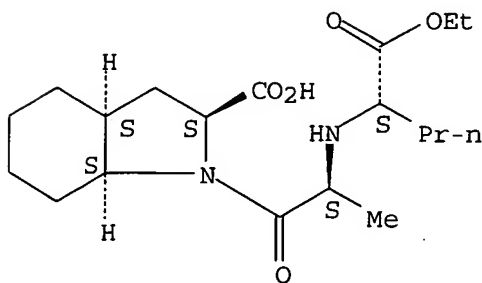
IT 82834-16-0P, Perindopril

RL: PNU (Preparation, unclassified); PREP (Preparation) (synthesis of [(ethoxycarbonyl)butyl]alanine for use in preparation of perindopril)

RN 82834-16-0 HCAPLUS

CN 1H-Indole-2-carboxylic acid, 1-[(2S)-2-[[[(1S)-1-(ethoxycarbonyl)butyl]amino]-1-oxopropyl]octahydro-, (2S,3aS,7aS)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).



REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS



## RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L20 ANSWER 7 OF 12 HCAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2004:266897 HCAPLUS

DOCUMENT NUMBER: 140:253917

TITLE: Process for the synthesis of  
perindopril and its pharmaceutically-  
acceptable salts

INVENTOR(S): Dubuffet, Thierry; Langlois, Pascal

PATENT ASSIGNEE(S): Les Laboratoires Servier, Fr.

SOURCE: Eur. Pat. Appl., 9 pp.

CODEN: EPXXDW

DOCUMENT TYPE: Patent

LANGUAGE: French

FAMILY ACC. NUM. COUNT: 1.

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 1403275	A1	20040331	EP 2003-290485	20030228
EP 1403275	B1	20051019		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, EL, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK				
AT 307139	E	20051115	AT 2003-290485	20030228
ES 2250846	T3	20060416	ES 2003-3290485	20030228
AU 2004217599	A1	20040916	AU 2004-217599	20040227
WO 2004078107	A2	20040916	WO 2004-FR446	20040227
WO 2004078107	A3	20041021		
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
CN 1753906	A	20060329	CN 2004-80005405	20040227
JP 2006519177	T2	20060824	JP 2006-500163	20040227
US 2006149081	A1	20060706	US 2005-547131	20050824
PRIORITY APPLN. INFO.:			EP 2003-290485	A 20030228
			WO 2004-FR446	A 20040227

OTHER SOURCE(S): MARPAT 140:253917

AB A method for the synthesis of perindopril involves coupling of (2S)-2,3,4,5,6,7-hexahydro-1H-indolecarboxylic acid (I) or an ester with N-[(S)-1-carbethoxybutyl]-L-alanine, followed by catalytic hydrogenation. I benzyl ester tosylate was prepared by reaction of 1-(1-cyclohexen-1-yl)pyrrolidine with (R)-ICH<sub>2</sub>CH(NBoc)CO<sub>2</sub>CH<sub>2</sub>Ph (Boc = tert-butoxycarbonyl), followed by deprotection and cyclization. Perindopril was converted into its tert-butylamine salt.

IT 82834-16-0P, Perindopril 107133-36-8P

RL: IMF (Industrial manufacture); SPN (Synthetic preparation); PREP (Preparation)

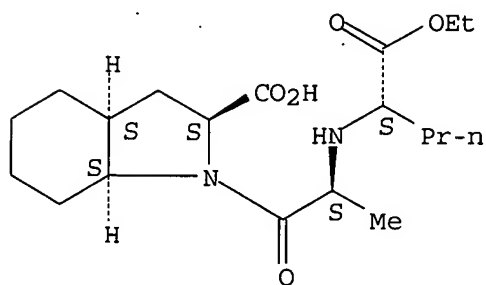
(synthesis of perindopril and pharmaceutically-acceptable salts)

RN 82834-16-0 HCAPLUS

CN 1H-Indole-2-carboxylic acid, 1-[(2S)-2-[(1S)-1-(ethoxycarbonyl)butyl]amino]-1-oxopropyl]octahydro-, (2S,3aS,7aS)-(9CI)  
(CA INDEX NAME)

10/11/2006 10566562a1.trn

Absolute stereochemistry. Rotation (-).



RN 107133-36-8 HCAPLUS

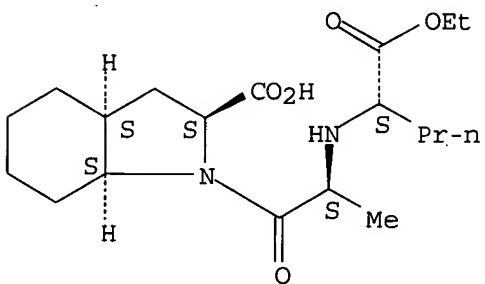
CN 1H-Indole-2-carboxylic acid, 1-[(2S)-2-[[[(1S)-1-(ethoxycarbonyl)butyl]amino]-1-oxopropyl]octahydro-, (2S,3aS,7aS)-, compd. with 2-methyl-2-propanamine (1:1) (9CI) (CA INDEX NAME)

CM 1

CRN 82834-16-0

CMF C19 H32 N2 O5

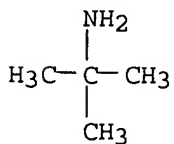
Absolute stereochemistry. Rotation (-).



CM 2

CRN 75-64-9

CMF C4 H11 N



REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L20 ANSWER 8 OF 12 HCAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2004:247009 HCAPLUS

DOCUMENT NUMBER: 140:253916

TITLE: Process for the synthesis of

N-[(S)-1-(ethoxycarbonyl)butyl]-(S)-alanine for use in preparation of perindopril  
 INVENTOR(S): Breard, Fabienne; Lecouve, Jean-Pierre  
 PATENT ASSIGNEE(S): Les Laboratoires Servier, Fr.  
 SOURCE: Eur. Pat. Appl., 9 pp.  
 CODEN: EPXXDW  
 DOCUMENT TYPE: Patent  
 LANGUAGE: French  
 FAMILY ACC. NUM. COUNT: 1  
 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 1400531	A1	20040324	EP 2003-292405	20030930
EP 1400531	B1	20060104		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, EE, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK				
AT 315046	E	20060215	AT 2003-292405	20030930
ES 2256693	T3	20060716	ES 2003-3292405	20030930
WO 2005033128	A1	20050414	WO 2004-FR2464	20040929
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW				
RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				

PRIORITY APPLN. INFO.: EP 2003-292405 A 20030930

OTHER SOURCE(S): MARPAT 140:253916

AB (S)-EtO2CCHPr-L-Ala-OH was prepared by a multistep procedure starting with allylation of Et glyoxylate with allylzinc bromide. Subsequent steps were resolution using Pseudomonas Fluorescens lipase, triflation of (R)-EtO2CCH(OH)CH2CH:CH2, substitution reaction with benzyl L-alaninate, and catalytic hydrogenolysis.

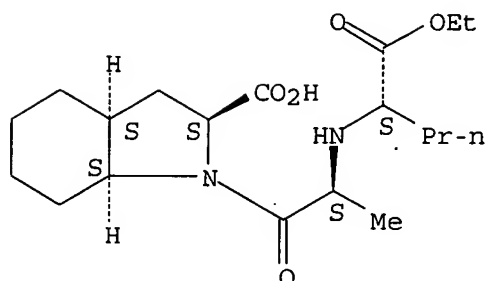
IT 82834-16-0P, Perindopril

RL: PNU (Preparation, unclassified); PREP (Preparation) (process for synthesis of (carbethoxybutyl)-L-alanine in preparation of perindopril)

RN 82834-16-0 HCAPLUS

CN 1H-Indole-2-carboxylic acid, 1-[(2S)-2-[[[(1S)-1-(ethoxycarbonyl)butyl]amino]-1-oxopropyl]octahydro-, (2S,3aS,7aS)-(9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).



REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L20 ANSWER 9 OF 12 HCAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2003:947713 HCAPLUS

DOCUMENT NUMBER: 139:381760

TITLE: Method for synthesis of perindopril and its pharmaceutically acceptable salts

INVENTOR(S): Dubuffet, Thierry; Lecouve, Jean-Pierre

PATENT ASSIGNEE(S): Les Laboratoires Servier, Fr.

SOURCE: Eur. Pat. Appl., 8 pp.

CODEN: EPXXDW

DOCUMENT TYPE: Patent

LANGUAGE: French

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 1367061	A1	<del>20031203</del>	EP 2003-291601	20030630
EP 1367061	B1	20060104		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK				
AT 315043	E	20060215	AT 2003-291601	20030630
ES 2256689	T3	20060716	ES 2003-3291601	20030630
AU 2004253721	A1	20050113	AU 2004-253721	20040628
WO 2005003153	A1	20050113	WO 2004-FR1637	20040628
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW				
RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG				
CN 1802384	A	20060712	CN 2004-80016014	20040628
US 2006178421	A1	20060810	US 2005-562490	20051222
PRIORITY APPLN. INFO.: EP 2003-291601 A 20030630				
WO 2004-FR1637 W 20040628				

OTHER SOURCE(S): CASREACT 139:381760; MARPAT 139:381760

AB A method for the synthesis of perindopril and its pharmaceutically-acceptable salts (e.g., the tert-butylamine) involves cyclocondensation reaction of N-[(S)-1-carbethoxybutyl]-(S)-alanine with

sulfinyl chlorides R1SOCl (R1 = imidazolyl, benimidazolyl, or tetrazolyl) to give Et (2S)-2-[(4S)-4-methyl-2,5-dioxo-1,2,3-oxathiazolidin-3-yl]pentanoate, which is amidated with (2S)-2,3,4,5,6,7-hexahydro-1H-indole-2-carboxylic acid and hydrogenated over 10% Pt/C to give perindopril.

IT 82834-16-0P, Perindopril 107133-36-8P

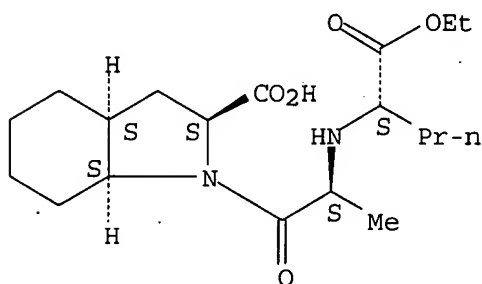
RL: IMF (Industrial manufacture); SPN (Synthetic preparation); PREP (Preparation)

(synthesis of perindopril via cyclocondensation of carbethoxybutylalanine with imidazolesulfinyl chloride)

RN 82834-16-0 HCAPLUS

CN 1H-Indole-2-carboxylic acid, 1-[(2S)-2-[(1S)-1-(ethoxycarbonyl)butyl]amino]-1-oxopropyl]octahydro-, (2S,3aS,7aS)- (9CI)  
(CA INDEX NAME)

Absolute stereochemistry. Rotation (-).



RN 107133-36-8 HCAPLUS

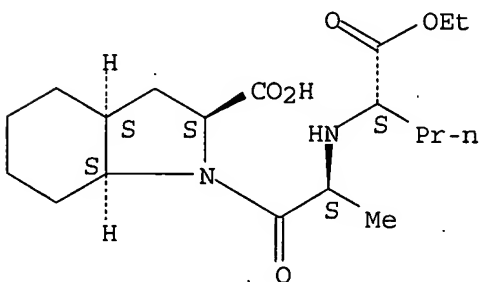
CN 1H-Indole-2-carboxylic acid, 1-[(2S)-2-[(1S)-1-(ethoxycarbonyl)butyl]amino]-1-oxopropyl]octahydro-, (2S,3aS,7aS)-, compd. with 2-methyl-2-propanamine (1:1) (9CI) (CA INDEX NAME)

CM . 1

CRN 82834-16-0

CMF C19 H32 N2 O5

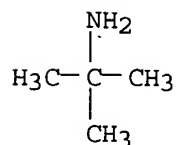
Absolute stereochemistry. Rotation (-).



CM 2

CRN 75-64-9

CMF C4 H11 N



REFERENCE COUNT: 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L20 ANSWER 10 OF 12 HCAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2003:910218 HCAPLUS

DOCUMENT NUMBER: 139:365227

TITLE: New process for the synthesis of N-[(S)-1-carboxybutyl]-(S)-alanine esters and their use in the synthesis of perindopril

INVENTOR(S): Breard, Fabienne; Fugier, Claude

PATENT ASSIGNEE(S): Les Laboratoires Servier, Fr.

SOURCE: Eur. Pat. Appl., 5 pp.

CODEN: EPXXDW

DOCUMENT TYPE: Patent

LANGUAGE: French

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

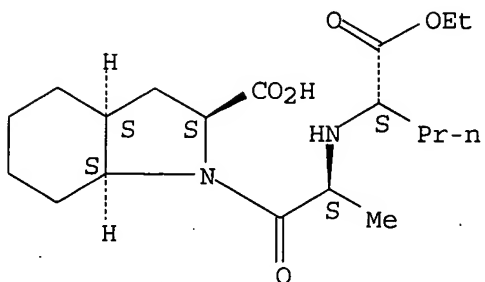
PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 1362845	A2	20031119	EP 2003-292145	20030901
EP 1362845	A3	20040331		
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK				
AU 2004270432	A1	20050317	AU 2004-270432	20040831
CA 2536926	AA	20050317	CA 2004-2536926	20040831
WO 2005023755	A1	20050317	WO 2004-FR2213	20040831
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW				
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CN 1835911	A	20060920	CN 2004-80023534	20040831
NO 2006001152	A	20060310	NO 2006-1152	20060310
PRIORITY APPLN. INFO.:			EP 2003-292145	A 20030901
			WO 2004-FR2213	W 20040831

OTHER SOURCE(S): CASREACT 139:365227; MARPAT 139:365227

AB Title alanine derivs. (S)-RO<sub>2</sub>CCHPr-L-Ala-OH (R = C<sub>1</sub>-C<sub>6</sub> alkyl) were prepared from N-protected (S)-5-methyl-2-morpholinone by propylation or allylation/hydrogenation, ring opening by LiOH, esterification, oxidation of the hydroxy group, and deprotection. In an example, N-[(S)-1-carbethoxybutyl]-(S)-alanine hydrochloride was prepared via allylation of Boc-protected (S)-5-methyl-2-morpholinone and treatment of tert-Bu (3S,5S)-5-methyl-3-propyl-2-oxo-4-morpholinecarboxylate with LiOH in aqueous MeCN and then EtI to afford intermediate Et (2S)-2-[(tert-

butoxycarbonyl)[(1S)-2-hydroxy-1-methylethyl]amino]pentanoate.  
 IT 82834-16-0P, Perindopril  
 RL: PNU (Preparation, unclassified); PREP (Preparation)  
 (process for synthesis of N-[(S)-carboxybutyl]-L-  
 alanine esters for use in synthesis of perindopril)  
 RN 82834-16-0 HCAPLUS  
 CN 1H-Indole-2-carboxylic acid, 1-[(2S)-2-[(1S)-1-  
 (ethoxycarbonyl)butyl]amino]-1-oxopropyl]octahydro-, (2S,3aS,7aS)-(9CI)  
 (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).



L20 ANSWER 11 OF 12 HCAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2003:609507 HCAPLUS

DOCUMENT NUMBER: 139:149930

TITLE: Process for the preparation of high purity  
perindopril and intermediates useful in its  
synthesis

INVENTOR(S): Simig, Gyula; Mezei, Tibor; Porcs-Makkay, Marta;  
Mandi, Attila

PATENT ASSIGNEE(S): Les Laboratoires Servier, Fr.

SOURCE: Eur. Pat. Appl., 12 pp.

CODEN: EPXXDW

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 1333026	A1	20030806	EP 2002-290206	20020130
R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, BG, MK, CY, AL, TR				
CA 2474003	AA	20030807	CA 2003-2474003	20030129
WO 2003064388	A2	20030807	WO 2003-IB691	20030129
WO 2003064388	A3	20040205		
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW				
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EE 200400107	A	20041015	EE 2004-107	20030129
BR 2003007293	A	20041221	BR 2003-7293	20030129
CN 1622936	A	20050601	CN 2003-802714	20030129
US 2005119492	A1	20050602	US 2003-503272	20030129
JP 2005521667	T2	20050721	JP 2003-564011	20030129
NO 2004003472	A	20040820	NO 2004-3472	20040820
BG 108858	A	20050531	BG 2004-108858	20040827

PRIORITY APPLN. INFO.:

EP 2002-290206	A	20020130
WO 2003-IB691	W	20030129

OTHER SOURCE(S): MARPAT 139:149930

AB The invention relates to 1-[2(S)-[1(S)-(ethoxycarbonyl)butylamino]propionyl]-[3a(S),7a(S)]octahydroindole-2(S)-carboxylic acid (perindopril) and its tert-butylamine salt, free of contaminants derivable from dicyclohexylcarbodiimide, and a process for their synthesis. The invention also relates to N-[1-(ethoxycarbonyl)butyl]-N-(alkoxycarbonyl)alanine intermediates used in the synthesis of perindopril, a known ACE inhibitor. Thus, N-[1-(ethoxycarbonyl)butyl]-N-(ethoxycarbonyl)alanine, prepared by ethoxycarbonylation of N-[1-(ethoxycarbonyl)butyl]alanine, was treated with thionyl chloride in CH<sub>2</sub>Cl<sub>2</sub> and acylated by perhydroindole-2-carboxylic acid in THF at reflux for 4-4.5 h. The product was treated with tert-butylamine to afford 55% perindopril ebumine.

IT 82834-16-0P, Perindopril 107133-36-8P,

Perindopril ebumine

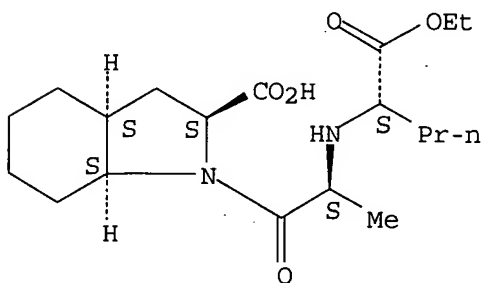
RL: IMF (Industrial manufacture); SPN (Synthetic preparation); PREP (Preparation)

(process for preparation of high purity perindopril and intermediates useful in its synthesis)

RN 82834-16-0 HCAPLUS

CN 1H-Indole-2-carboxylic acid, 1-[(2S)-2-[[[(1S)-1-(ethoxycarbonyl)butyl]amino]-1-oxopropyl]octahydro-, (2S,3aS,7aS)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).



RN 107133-36-8 HCAPLUS

CN 1H-Indole-2-carboxylic acid, 1-[(2S)-2-[[[(1S)-1-(ethoxycarbonyl)butyl]amino]-1-oxopropyl]octahydro-, (2S,3aS,7aS)-, compd. with 2-methyl-2-propanamine (1:1) (9CI) (CA INDEX NAME)

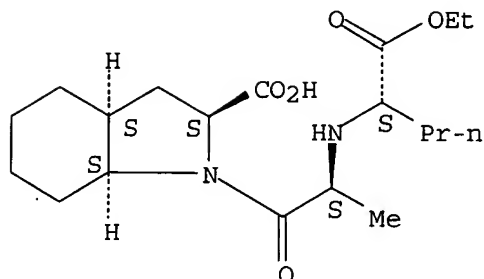
CM 1

CRN 82834-16-0

CMF C19 H32 N2 O5

Absolute stereochemistry. Rotation (-).

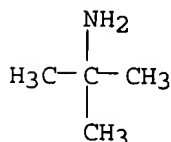




CM 2

CRN 75-64-9

CMF C4 H11 N



REFERENCE COUNT: 2 THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L20 ANSWER 12 OF 12 HCAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2002:641139 HCAPLUS

DOCUMENT NUMBER: 138:142607

TITLE: On-line simultaneous determination of S- and R-perindopril using amperometric biosensors as ~~detectors in flow systems~~

AUTHOR(S): Stefan, Raluca-Ioana; van Staden, Jacobus F.; Mulaudzi, Ludwig Vusimuzi; Aboul-Enein, Hassan Y.  
CORPORATE SOURCE: Department of Chemistry, University of Pretoria, Pretoria, 0002, S. Afr.

SOURCE: Analytica Chimica Acta (2002), 467(1-2), 189-195  
CODEN: ACACAM; ISSN: 0003-2670

PUBLISHER: Elsevier Science B.V.

DOCUMENT TYPE: Journal

LANGUAGE: English

AB Two types of flow systems were selected for the simultaneous assay of S- and R-perindopril (pdp): flow injection anal. (FIA) and sequential injection anal. (SIA). The SIA system was more efficient, because of the highest precision and accuracy, and the lower consumption of sample and buffer. The amperometric biosensors used as detectors in the flow systems were based on L- and D-amino acid oxidase (AAOD). The linear concentration ranges are in the nmol L<sup>-1</sup> range, from 120 pmol L<sup>-1</sup> to 40 nmol L<sup>-1</sup> (3+S.D.), with very low detection limits. The biosensors/flow system can be used reliably online in synthesis process control, for the simultaneous assay of S- and R-pdp with a frequency of more than 30 samples per h.

IT 82834-16-0, Perindopril 145513-48-0

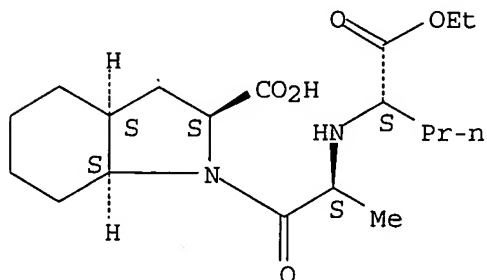
RL: ANT (Analyte); ANST (Analytical study).  
(resolution of perindopril using amperometric biosensors as detectors in flow systems)

10/11/2006 10566562a1.trn

RN 82834-16-0 HCAPLUS

CN 1H-Indole-2-carboxylic acid, 1-[(2S)-2-[[[(1S)-1-(ethoxycarbonyl)butyl]amino]-1-oxopropyl]octahydro-, (2S,3aS,7aS)-(9CI)  
(CA INDEX NAME)

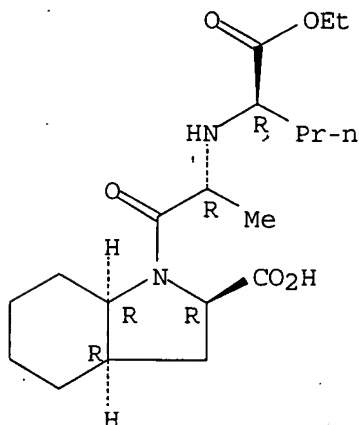
Absolute stereochemistry. Rotation (-).



RN 145513-48-0 HCAPLUS

CN 1H-Indole-2-carboxylic acid, 1-[(2R)-2-[[[(1R)-1-(ethoxycarbonyl)butyl]amino]-1-oxopropyl]octahydro-, (2R,3aR,7aR)-(9CI)  
(CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT:

22

THERE ARE 22 CITED REFERENCES AVAILABLE FOR THIS  
RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

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COST IN U.S. DOLLARS

SINCE FILE	TOTAL
ENTRY	SESSION
73.97	931.15

FULL ESTIMATED COST

DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS)

SINCE FILE	TOTAL
ENTRY	SESSION
-9.00	-9.75

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STN INTERNATIONAL LOGOFF AT 13:44:03 ON 11 OCT 2006